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From the INTERNATIONAL SEARCHING AUTHORITY	PCT			
To:	NOTIFICATION OF TRANSMITTAL OF			
GLAXOSMITHKLINE	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT AND			
Attn. Levy, David J. TERRET VIEW Five Moore Drive. P.O. DBOX NO. 18.	THE WRITTEN OPINION OF THE INTERNATIONAL			
	() (SEARCHING AUTHORITY, OR THE DECLARATION			
Research Triangle Park (NC 27709 UNITED STATES OF AMERICA: MAR 0 4 2005				
MAR 0 4 2003				
<u></u>	(PCT Rule 44.1)			
GLOBAL INTELLECTUAL PRO	PRIV			
	(day/month/year) 24/02/2005			
Applicant's or agent's file reference	<u> </u>			
PU4831WO	FOR FURTHER ACTION See paragraphs 1 and 4 below			
International application No.	International filing date			
PCT/US2004/018456	(day/month/year) 09/06/2004			
Applicant	:			
SMITHKLINE BEECHAM CORPORATION				
The applicant is hereby notified that the international search Authority have been established and are transmitted herewi	report and the written opinion of the International Searching			
Filing of amendments and statement under Article 19:	to the transfer of the Property of the ACA			
The applicant is entitled, if he so wishes, to amend the claim When? The time limit for filing such amendments is non	ns of the International Application (see Hule 45):			
International Search Report; however, for more	details, see the notes on the accompanying sheet.			
Where? Directly to the International Bureau of WIPO, 34 1211 Geneva 20, Switzerland, Fa	chemin des Colombettes ascimile No.: (41–22) 740.14.35			
For more detailed instructions, see the notes on the acco				
2. The applicant is hereby notified that no international search Article 17(2)(a) to that effect and the written opinion of the Ir	report will be established and that the declaration under iternational Searching Authority are transmitted herewith.			
3. With regard to the protest against payment of (an) addition	onal fee(s) under Rule 40.2, the applicant is notified that:			
the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the texts of both the protest and the decision thereon to the designated Offices.				
no decision has been made yet on the protest; the app	olicant will be notified as soon as a decision is made.			
4. Reminders				
Shortly after the expiration of 16 months from the priority date, the International Bureau. If the applicant wishes to avoid or postpone application, or of the priority claim, must reach the International B before the completion of the technical preparations for internation	publication, a notice of withdrawal of the international ureau as provided in Rules 90 <i>bls</i> .1 and 90 <i>bls</i> .3, respectively, nal publication.			
The applicant may submit comments on an informal basis on the International Bureau. The International Bureau will send a copy o international preliminary examination report has been or is to be a the public but not before the expiration of 30 months from the price	f such comments to all designated Onices unless all established. These comments would also be made available to			

Within 19 months from the priority date, but only in respect of some designated Offices, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later); otherwise, the applicant must, within 20 months from the priority date, perform the prescribed acts for entry into the national phase before those designated Offices.

In respect of other designated Offices, the time limit of **30 months** (or later) will apply even if no demand is filed within 19 months.

See the Annex to Form PCT/IB/301 and, for details about the applicable time limits, Office by Office, see the PCT Applicant's Guide, Volume II, National Chapters and the WIPO Internet site.

Name and n	nailing address of the International Searching Authority
<u></u>	European Patent Office, P.B. 5818 Patentiaan 2

NL-2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016 Authorizad officer

Angela Lopez Navarro



INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference PU4831WO	FOR FURTHER ACTION		see Form PCT/ISA/220 ;, where applicable, item 5 below.
International application No.	International filing date (day/monti	h/year)	(Earliest) Priority Date (day/month/year)
			10/05/0000
PCT/US2004/018456	09/06/2004		10/06/2003
Applicant			
SMITHKLINE BEECHAM CORPOR	ATION		
This International Search Report has bee according to Article 18. A copy is being to	n prepared by this international Sea ansmitted to the international Bureau	rching Author J.	ity and is transmitted to the applicant
This International Search Report consists	of a total of 11 she	ets.	<i>:</i>
X It is also accompanied by	a copy of each prior art document c	ited in this rep	port.
	·		
Basis of the report With regard to the language, the language in which it was filed, un	international search was carried out less otherwise indicated under this it	on the basis em.	of the International application in the
The international this Authority (Ru		of a translatio	on of the International application furnished to
b. With regard to any nucle	otide and/or amino acid sequence	disclosed in	the international application, see Box No. I.
2. X Certain claims were fou	nd unsearchable (See Box II).		`:
3. Unity of invention is lac	king (see Box III).		··· .
4. With regard to the title,	ubmitted by the applicant.		
	shed by this Authority to read as folio	ws:	
			COCORTICOID, MINERALOCORTICOID
			•
AND PROGESTERONE RECEI	PTORS		.`
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			·
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5. With regard to the abstract,	at anything of the address of the manufacture is		
· ·	ubmitted by the applicant.	hie Authofh/	as It appears in Box No. IV. The applicant
may, within one month for	om the date of mailing of this interna	tional search	report, submit comments to this Authority.
6. With regard to the drawings,			
a. the figure of the drawings to be	published with the abstract is Figure	No	:
as suggested by			
,	is Authority, because the applicant for	ailed to sugge	est a ligure.
[is Authority, because this figure bett		
	be published with the abstract.		· ·
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Form PCT/ISA/210 (first sheet) (January 2004)

PCT/US2004/018456 A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C07C211/59 C07C C07C211/60 C07D295/13 C07C215/16 C07C229/68 A61K31/04 A61K31/136 A61K31/277 A61K31/4453 A61P5/26 According to International Patent Classification (IPC) or to both national classification and IPC Minimum documentation searched (classification system followed by classification symbols) IPC 7 C07C C07D Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the International search (name of data base and, where practical, search terms used) EPO-Internal, CHEM ABS Data, BEILSTEIN Data C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to daim No. Citation of document, with indication, where appropriate, of the relevant passages Category * 1,3,6, X DATABASE BEILSTEIN 10-13 BEILSTEIN INSTITUTE FOR ORGANIC CHEMISTRY, FRANKFURT-MAIN, DE: XP002312536 retrieved from XFIRE Database accession no. 9508573 abstract & MOLECULES, vol. 7, no. 12, 2002, pages 885-895, 1,3,6,7, X DATABASE BEILSTEIN BEILSTEIN INSTITUTE FOR ORGANIC CHEMISTRY. 10-13 FRANKFURT-MAIN, DE; XP002312537 retrieved from XFIRE Database accession no. 8221473 abstract _/---Further documents are listed in the continuation of box C. Patent family members are listed in annex. · Χl Special categories of cited documents: T later document published after the International filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive slep when the document is taken alone "E" earlier document but published on or after the International filing date 'L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled "O" document referring to an oral disclosure, use, exhibition or

- "P" document published prior to the international filing date but later than the priority date claimed
- "&" document member of the same patent family

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Date of the actual completion of the international search

7. January 2005

24/02/2005

7 January 2005

Name and mailing address of the ISA

European Patent Office, P.B. 5616 Patentiaan 2

Ni. – 2280 HV Rijswijk

Tel. (+31–70) 340–2040. Tx. 31 651 epo nl.

Fax: (+31–70) 340–3016

Authorized officer

Cooper, S

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	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	•
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	vol. 46, no. 8, 1998, pages 1265-1273,	≎ `
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	FRANKFURT-MAIN, DE;	•
	XP002312538	
	retrieved from XFIRE	·
	Database accession no. 7464824	
	abstract & J. MED. CHEM.,	•
	vol. 38, no. 26, 1995, pages 5051-5065,	
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	BEILSTEIN INSTITUTE FOR ORGANIC CHEMISTRY,	10-13
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	abstract	
	& J. MED. CHEM., vol. 38, no. 8, 1995, pages 1344-1354,	5.
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x l	J. DONALD ALBRIGHT ET AL: "Potential	1-3,6,7,
1	Antiatherosclerotic Agents. 3. Substituted	10-13
	Benzoic and Non Benzoic Acid Analogues of	Ì
	Cetaben"	
	J. MED. CHEM,	:
	vol. 26, no. 10, 1983, pages 1393-1411,	· ·
	XP002312531	
	Scheme V; Table IV, compound 84	:
x l	DATABASE BEILSTEIN	1,3,
"	BEILSTEIN INSTITUTE FOR ORGANIC CHEMISTRY,	10-13
	FRANKFURT-MAIN, DE;	· ·
	XP002312540	·
	retrieved from XFIRE	
	Database accession no. 5747353, 3281950	· · · · · · · · · · · · · · · · · · ·
	abstract	<u> </u>
	& CHEM. PHARM. BULL.,	
	vol. 32, no. 10, 1984, pages 3968-3980,	· ·
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	abstract	l see
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X	YI-JEN SHUE ET AL: "Direct palladium(0)-catalyzed amination of allylic alcohols with aminonaphthalenes" TETRAHEDRON LETTERS, vol. 44, no. 7,	1,2,7
	10 February 2003 (2003-02-10), pages 1481-1485, XP004405249 Table 2, compounds 1c,3c,4c,1d,3d,4d,1f,3f	
X	IRINA A. BALOVA ET AL: "A one-pot synthesis of 1-arylalka-1,3-diynes by sequential acetylene zipper and Sonogashira reactions"	1-3
	TETRÄHEDRON LETTERS, vol. 44, no. 1, 1 January 2003 (2003-01-01), pages 107-109, XP004397130 Page 108: compounds 7a and 8a and starting aryl iodide	
K	YI XIAO AND XUHONG QIAN: "Novel highly efficient fluoroionophores with a peri-effect and strong electron-donating receptors: TICT-promoted PET and signaling response to transition metal cations with low background emission" TETRAHEDRON LETTERS,	1-3,6,7
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	vol. 43, no. 16, 15 April 2002 (2002-04-15), pages 2991-2994, XP004345946 Page 2992: Compounds E, H1-H4,G	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\
X	EDWARD F. ELSLAGER ET AL: "Synthetic Schistosomicides. VI. 4-Substituted 1-(Dialkylaminoalkylamino)naphthalenes" J. MED. CHEM., vol. 7, 1964, pages 658-662, XP002312532 Page 660, Table I; Chart II, compound VIII; compounds XVIIIa,b	1-3,5-7
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Calegory •	etion) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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	abstract	
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	BEILSTEIN INSTITUTE FOR ORGANIC CHEMISTRY,	,
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	J. MED. CHEM.,	
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	VEGYESZET) 14 May 1970 (1970-05-14)	1
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Category *	cition) DOCUMENTS CONSIDERED TO BE RELEVANT	Debuggita dela si
Caregory -	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
х	DATABASE BEILSTEIN BEILSTEIN INSTITUTE FOR ORGANIC CHEMISTRY, FRANKFURT-MAIN, DE;	1,2
	XP002312546 retrieved from XFIRE Database accession no. 641096 abstract	
-	& J. ORG. CHEM., vol. 37, 1972, pages 3248-3252,	•
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	retrieved from XFIRE Database accession no. 409369 abstract & J. MED. CHEM.,	·
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	FRANKFURT-MAIN, DE; XP002312548 retrieved from XFIRE Database accession no. 318879	
	abstract & J. PRAKT. CHEM., vol. 60, no. 2, 1899, page 193,	:
(DATABASE BEILSTEIN BEILSTEIN INSTITUTE FOR ORGANIC CHEMISTRY, FRANKFURT-MAIN, DE; XP002312549 retrieved from XFIRE	1,2
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WO 03/011824 A (SQUIBB BRISTOL MYERS CO; SUN CHONGQING (US); AUGERI DAVID (US); HAMAN) 13 February 2003 (2003-02-13) Page 50, line 24; page 55, lline 8; page 62, line 4; page 63, line 21 - page 64, line 2.	1-3
claims 1-16	14-17
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page 1, lines 5-7; claims 1-8	14-17
WO 02/24702 A (BALOG JAMES AARON; PICKERING DACIA A (US); SQUIBB BRISTOL MYERS CO (U) 28 March 2002 (2002-03-28) Page 173, line 16 - page 174, line 7;	1-3
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G. BARTOLI ET AL: "Reduction of Nitronate Adducts from RMgX and 1-Methoxy-4-nitro-naphthalene to Oximes or Amines by Trivalent Phosphorus Reagents" TETRAHEDRON, vol. 40, no. 18, 1984, pages 3437-3442, XP002312534 Page 3438: Compounds 6a-c,6e	1
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PCT/US2004/018456

	MON) DOCUMENTS CONSIDERED TO BE RELEVANT	·
egory °	Cliation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	NIGEL J. BUNCE AND STEPHEN R. CATER: "Photosubstitution of 1-Methoxy-4-nitronaphthalene with Amine Nucleophiles: Dual Pathways" J. ORG. CHEM., vol. 52, 1987, pages 4214-4223, XP002312535 Scheme I: Compounds 2 on p.4216: compounds	1-3
	Scheme I; Compounds 2 on p.4216; compounds 3 on p.4217; Experimental section	·
		<u>;</u>
		·
		.46
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INTERNATIONAL SEARCH REPORT

PC1/U52004/018450

Box II Observations where certain claims were found unsearchable (Continuation of item 2 of fire	st sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following	owing reasons:
1. X Claims Nos.:	•
Although claims 16 and 17 are directed to a method of the human/animal body, the search has been carried out and ba alleged effects of the compound/composition.	of treatment sed on the
2. Claims Nos.:	- ₹. - ₹ - .*
because they relate to parts of the International Application that do not comply with the prescribed requireme an extent that no meaningful International Search can be carried out, specifically:	nts to such
mi data in meningar meneral da ana ana ana ana ana ana ana ana ana	•
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□ g ₁₋₁	•
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of	f <u>Fule 6.4(a)</u> .
Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)	
and in the second multiple inventions in this interest and institute as follows:	
This international Searching Authority found multiple inventions in this international application, as follows:	÷
	;
1. As all required additional search fees were timely paid by the applicant, this International Search Report cov	eue a n
searchable daims.	
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not in of any additional fee.	vite payment
3. As only some of the required additional search fees were timely paid by the applicant, this international Sea covers only those claims for which fees were paid, specifically claims Nos.:	rch Report
4. No required additional search fees were timely paid by the applicant. Consequently, this international Search restricted to the invention first mentioned in the claims; it is covered by claims Nos.:	h Report is
	•
Remark on Protest The additional search fees were accompanied by the a	oplicant's protest.
•••••••••••••••••••••••••••••••••••••	
No protest accompanied the payment of additional sea	rch fees.

PCT/US2004/018456

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NOTES TO FORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the fifing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published to the purposes of provisional protection or has another reason for amending the claims before international polication. Furthermore, it should be emphasized that provisional protection is available in some "states only."

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When?

Within 2 months from the date of transmittel of the international search report or 16 months from the priority date, whichever time timit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Pule 46.1).

Where not to file the smendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been its filed, see below.

How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The emendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims, it should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

NOTES TO FORM PCT/ISA/220 (continued)

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- [Where originally there were 48 claims and after amendment of some claims there are 51]:
 "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers;
 claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- (Where originally there were 14 claims and the amendments consist in carcelling some claims and in adding new claims):
 Claims 1 to 5 and 14 constructed claims 7 to 13 approximate year desires 15, 16 and 17 added. or
 - "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- [Where various kinds of amendments are made]:
 "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claims 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the latter indicating the differences between the claims as filed and as amended. It must be filed on a separate aheat and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62-2(a), first sentence).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide.